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<u>Preosteoblast-enriched lnc-Evf2 facilitates osteogenic differentiation by targeting</u> <u>Notch</u>

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Ossification of ligaments (OL) and <u>osteoporosis</u> (OP) are multifactorial disorders without definitive clinical biomarkers. Long non-coding RNAs (lncRNAs) are known to involve in regulating pathogenesis. Here, we have identified a preosteoblast-enriched lnc-Evf2 that was overexpressed in <u>ossified ligamentum flavum</u> (OLF) and down-expressed in OP. lnc-Evf2 is gradually upregulated during osteogenic induction, correlating with the enhanced expression of osteogenic marker genes and matrix mineralization. Moreover, knockdown of lnc-Evf2 significantly inhibits the expression of osteogenic differentiation markers and delays the osteoblastic mineralization process, indicating that this molecule is involved in <u>osteogenesis</u>. Mechanistically, we demonstrated that silencing of lnc-Evf2 decreases the protein level but not the mRNA levels of Notch2, Notch3, and Hes1, all of which correlate with osteogenesis. Taken together, our data demonstrate that lnc-Evf2 promotes osteogenic differentiation and bone formation through the Notch signaling, revealing that lnc-Evf2 may serve as a novel potential clinical target of OL and OP.



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Recent Publications

- Harper Zhen Zhang, Haixia Q, Han Xia#, Qi Liu, Yi Ren, Kun Zhang, Yuan Xue, and Wei Hong1. Preosteoblast-enriched lnc-Evf2 facilitates osteogenic differentiation by targeting Notch. Acta *Biochim Biophys Sin* (Shanghai). 2021 Feb 4;53(2):179-188.
- Yutao Tang, Han Xia, et al. Effects of Intermittent Parathyroid Hormone 1-34 Administration on Circulating Mesenchymal Stem Cells in Postmenopausal Osteoporotic Women. *Med Sci Monit*. 2019 Jan 8; 25:259-268.
- 3. Yawei Han, Kun Zhang, Yuheng Hong, Jingzhao Wang, Qi Liu, Zhen Zhang, Han Xia, Yutao Tang, Tengshuai Li, Liandong Li, Yuan Xue, Wei Hong. miR-342-3p promotes osteogenic differentiation via targeting ATF3. *FEBS Lett.* 2018 Dec;592(24):4051-4065.
- 4. Yawei Han, Yuheng Hong, Liandong Li, Tengshuai Li, Zhen Zhang, Jingzhao Wang, Han Xia, Yutao Tang, Zhemin Shi, Xiaohui Han, Ting Chen, Qi Liu, Mengxia Zhang, Kun Zhang, Wei Hong, Yuan Xue. A Transcriptome-Level Study Identifies Changing Expression Profiles for Ossification of the Ligamentum Flavum of the Spine. *Mol Ther Nucleic Acids*. 2018 Sep 7; 12:872-883.
- 5. Jiaming Zhou, et al. Clinical efficacy of calcitonin compared to diclofenac sodium in chronic nonspecific low back pain with type I Modic changes: a retrospective study. *J Pain Res.* 2018 Jul 17;11:1335-1342.

Biography

Han Xia has his expertise in evaluation and passion in improving the health and wellbeing. In the current study, he showed that Inc-Evf2 is a preosteoblast-enriched IncRNA that is overexpressed in OLF and during osteogenic differentiation, but was down-expressed in OP. Knockdown of Inc-Evf2 significantly inhibits the expression of osteogenic differentiation marker genes and the ALP activity and delays the osteoblastic mineralization process. Mechanistically, we demonstrated that silencing of Inc-Evf2 decreases the protein levels but not the mRNA levels of Notch2, Notch3, and Hes1 in preosteoblasts and mouse embryonic fibroblasts, suggesting that Inc-Evf2 promotes <u>osteogenesis</u> via the Notch signaling. focusing.

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